

FIGURE 1

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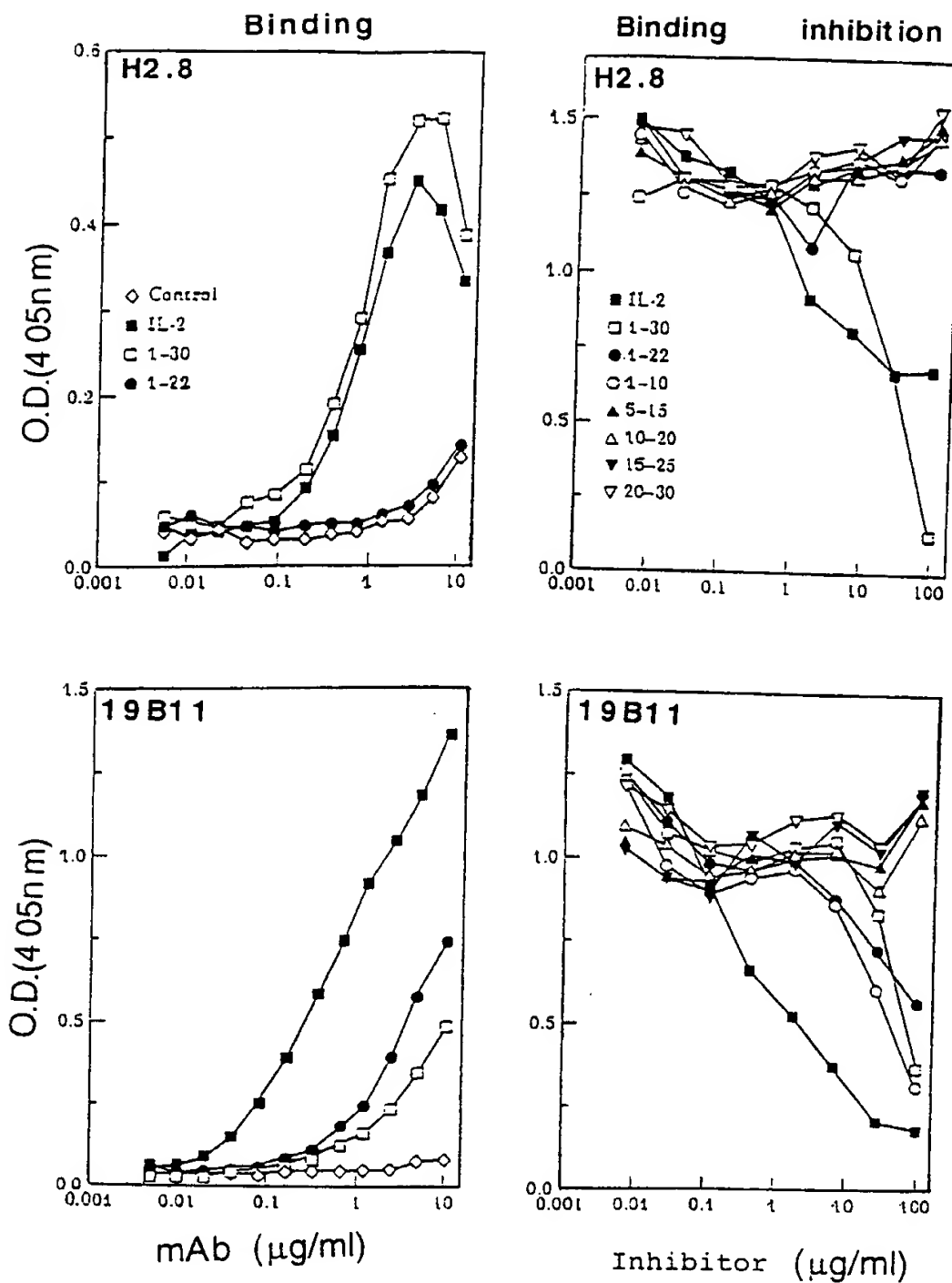


FIGURE 2

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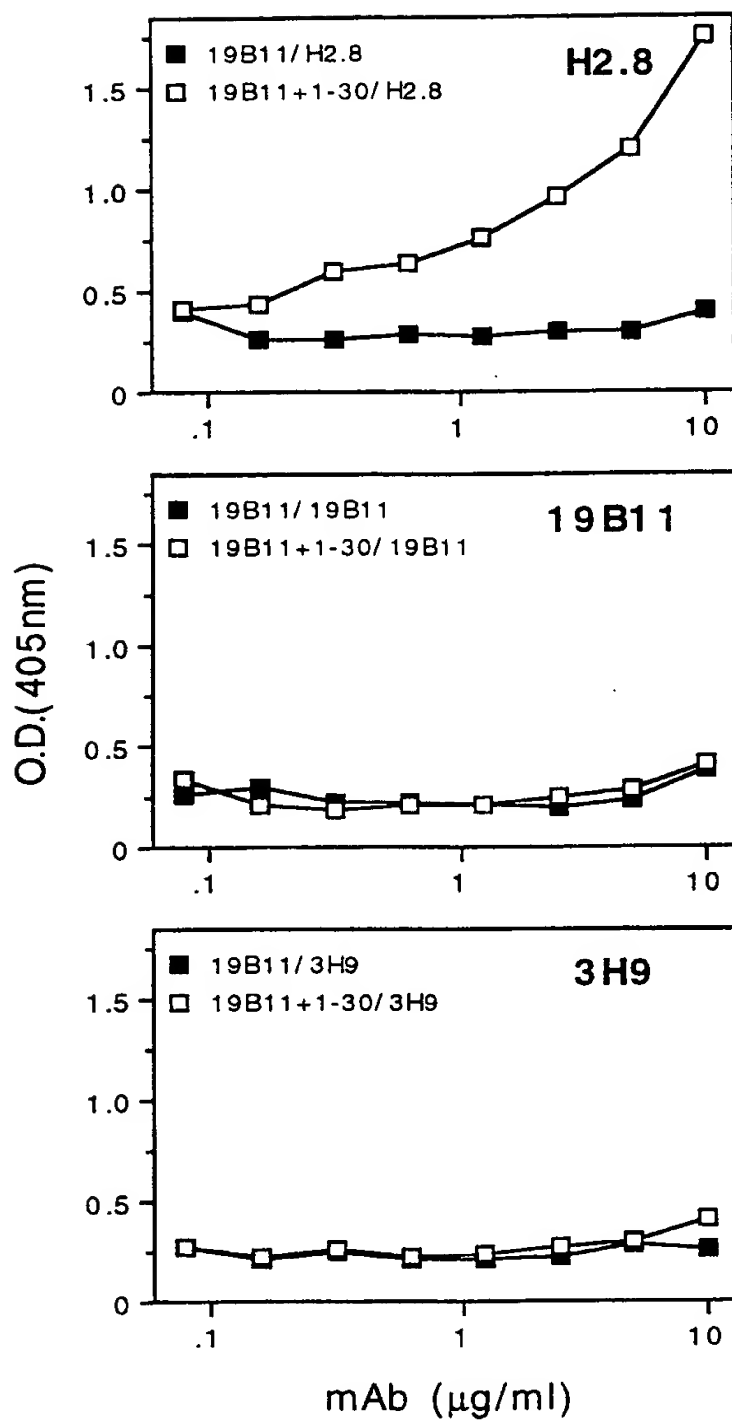


FIGURE 3

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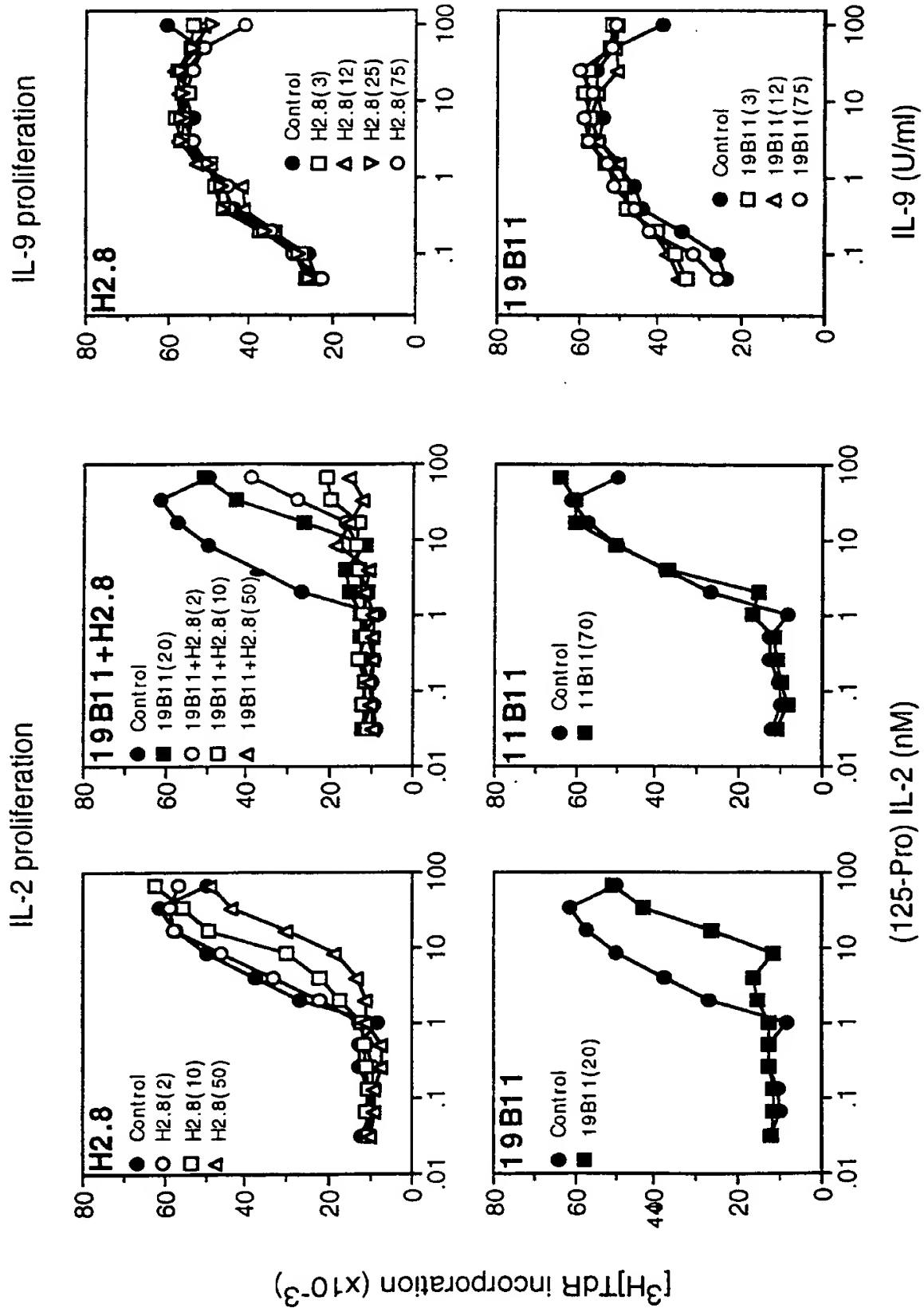


FIGURE 4

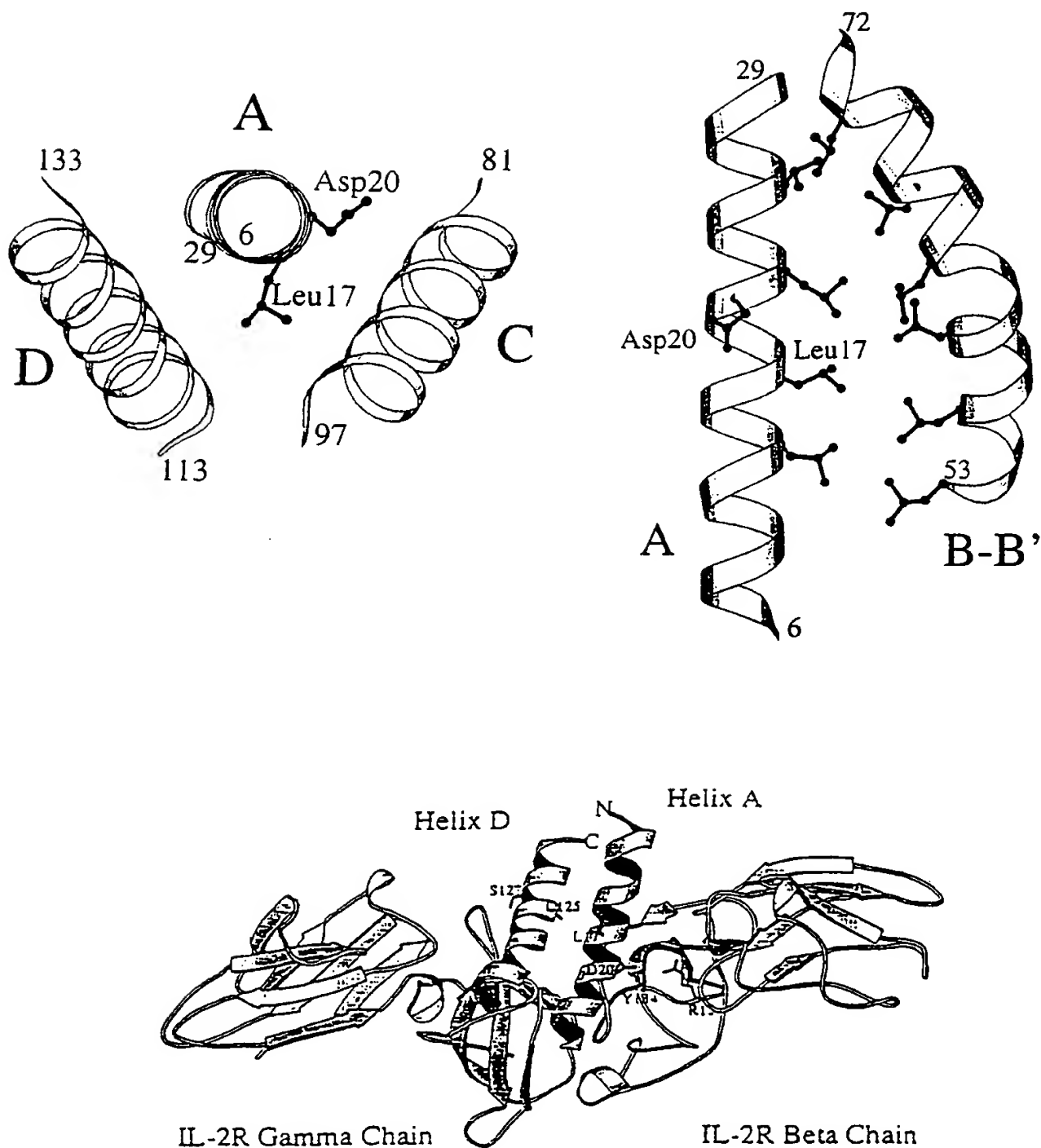
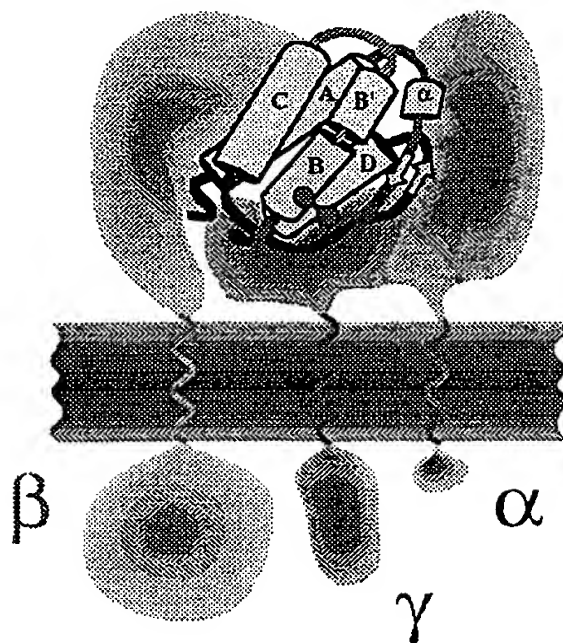


FIGURE 5

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Interleukine-2 receptor



IL-2 AND IP130 SEQUENCE (α -HELICES ARE BOXED)

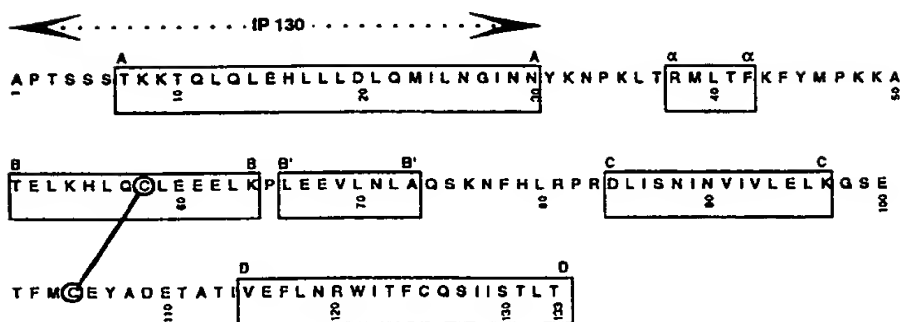


FIGURE 6

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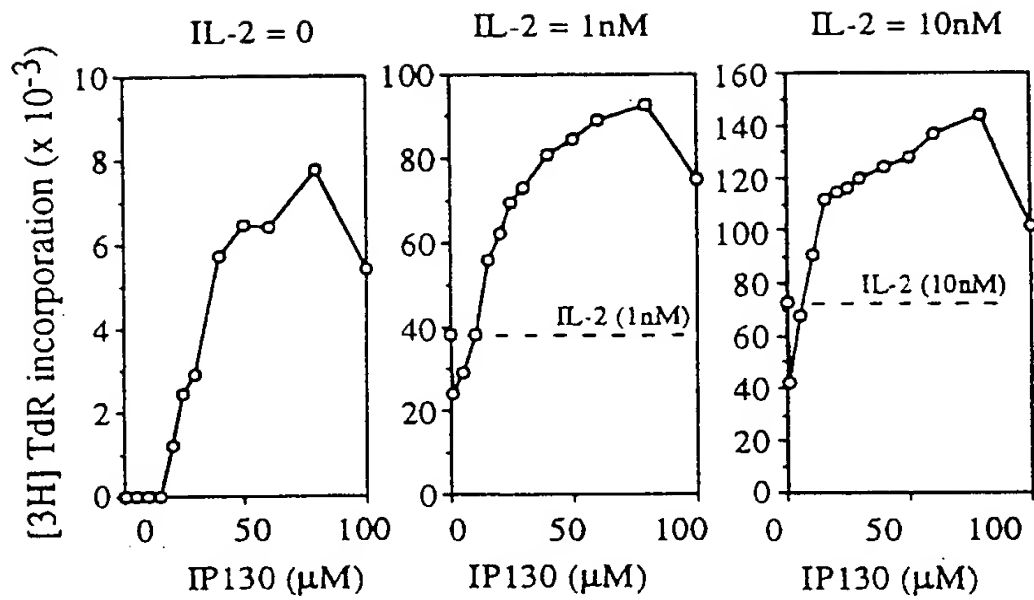


FIGURE 7

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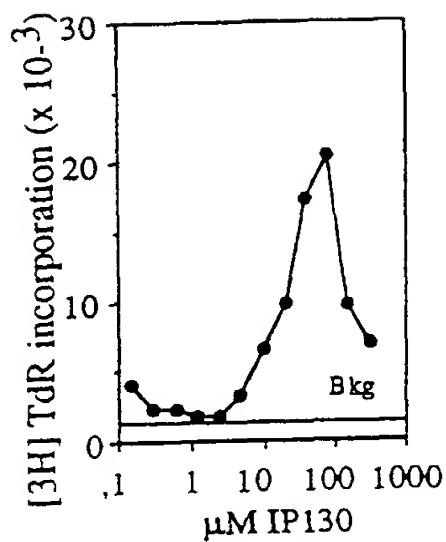


FIGURE 8A

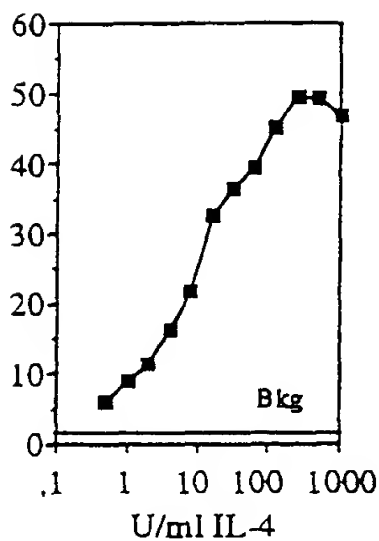


FIGURE 8B

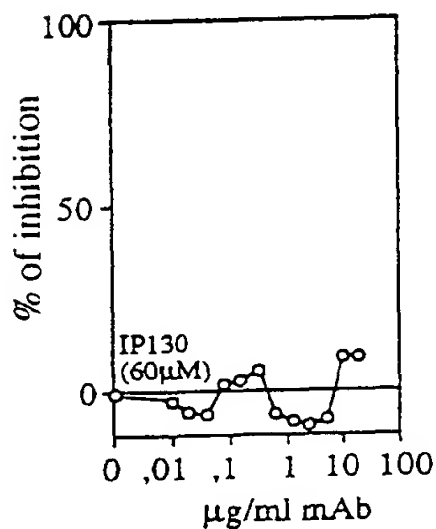


FIGURE 8C

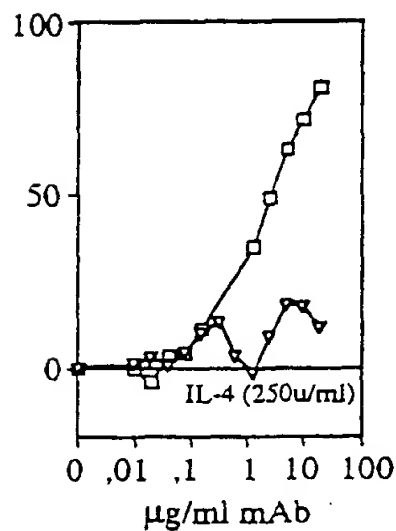


FIGURE 8D

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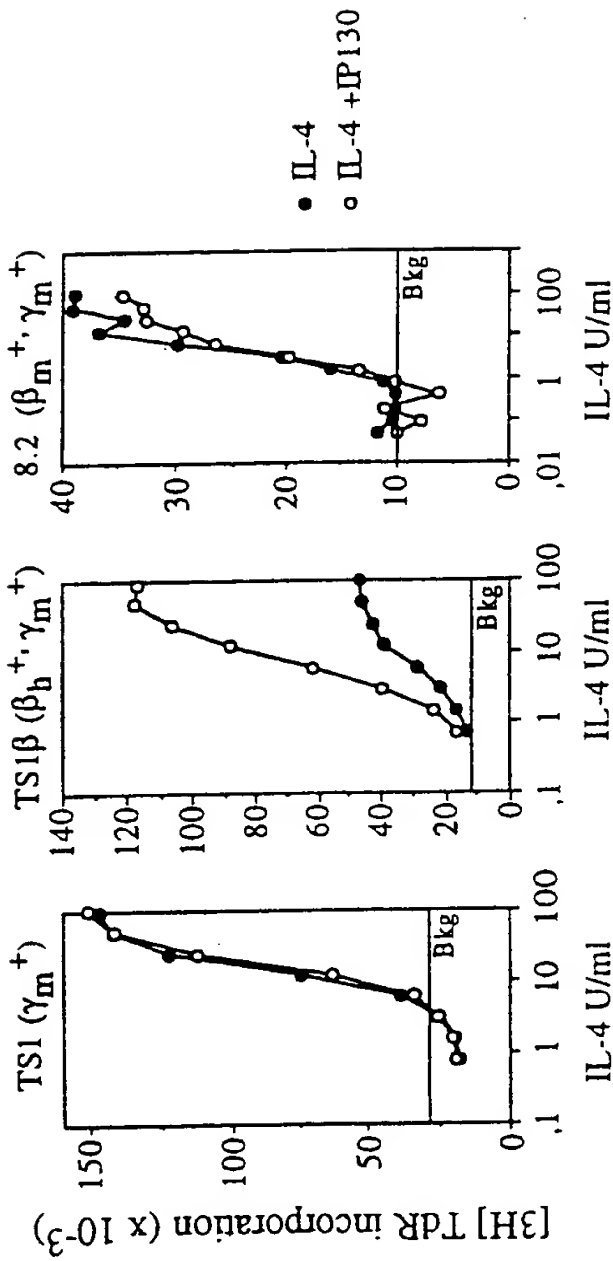


FIGURE 9A

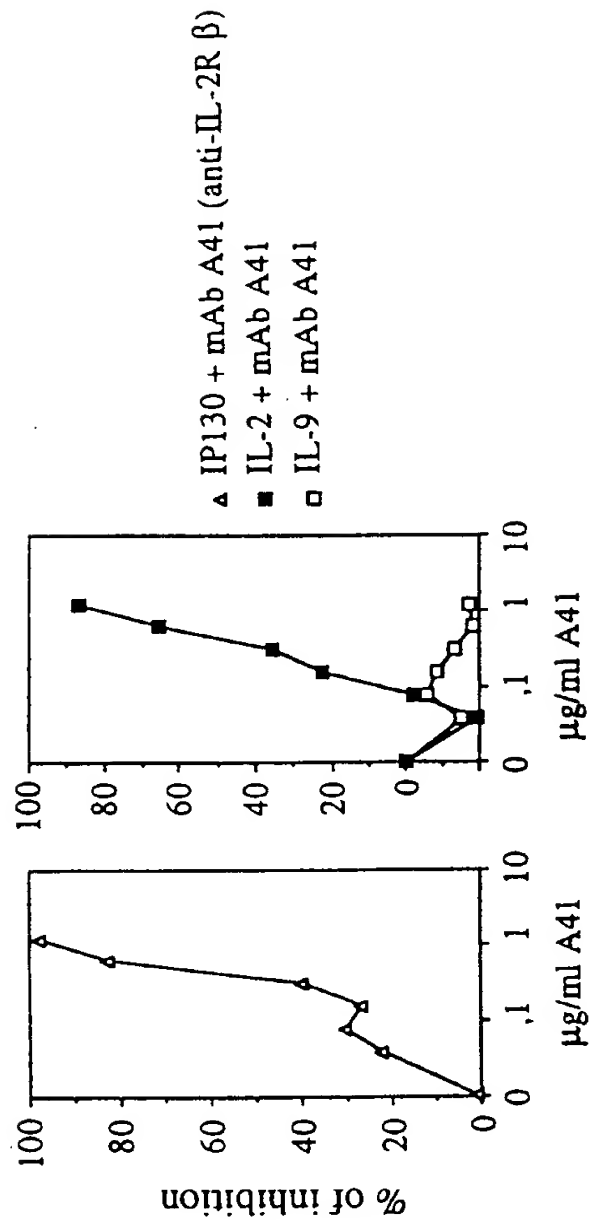
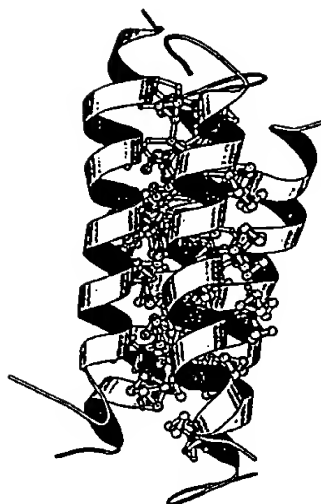


FIGURE 9B

1 APTTSSSTKKKTQLQLLEHLLDLQMLNGINN	10 20 30	% helix (Circular dichroism)	Main molecular species	Activity
1	30	50% (150 Å 30µM) 35% (4µM)	Tetramer (4M-8M, Kd=30-100µM) /Octamer	+++
1	30	22% (150 Å 30µM)	Dimer (1M-2M, Kd=0.2µM) /Tetramer (2M-4M, Kd=100µM)	++
1	22	<2%		-
1	10	0%	Dimer (1M-2M, Kd=50µM) (2M-4M, Kd=1.4mM)	-
5	15	0%	Dimer (1M-2m, Kd=113µM)	-
1	20	0%	Monomer	ND
1	30	<5%	Monomer	+

FIGURE 10

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IP130

IP130

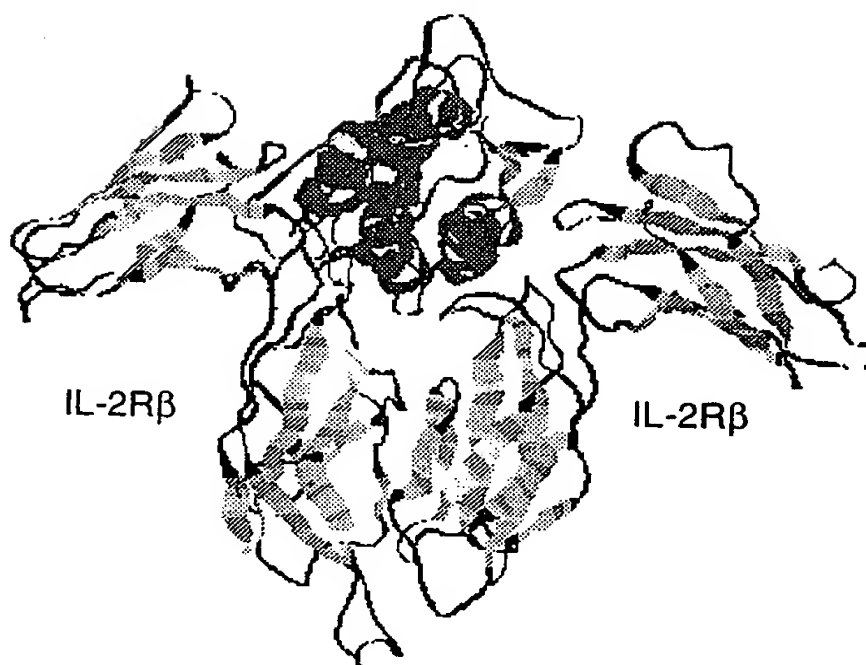
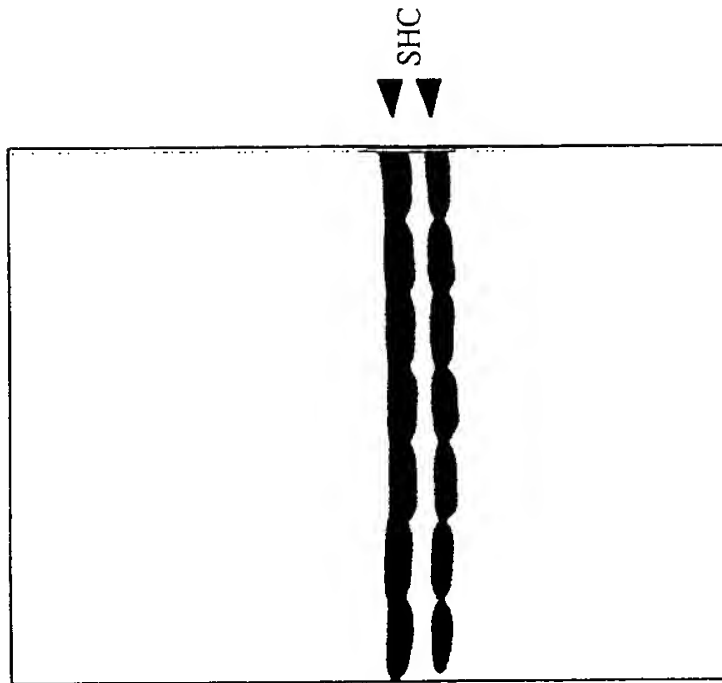


FIGURE 11

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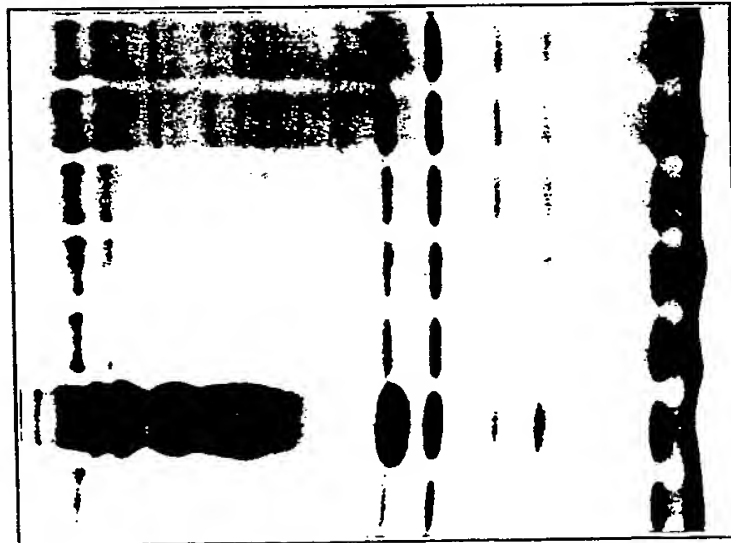
BLOT ANTI SHC



no stimulation
IL-2 (10 min)
IP130 (1 min)
IP130 (2 min)
IP130 (5 min)
IP130 (10 min)
IP130 (15 min)

FIGURE 12B

BLOT 4G10 (ANTI
PHOSPHOTYROSINE)



no stimulation
IL-2 (10 min)
IP130 (1 min)
IP130 (2 min)
IP130 (5 min)
IP130 (10 min)
IP130 (15 min)

FIGURE 12A

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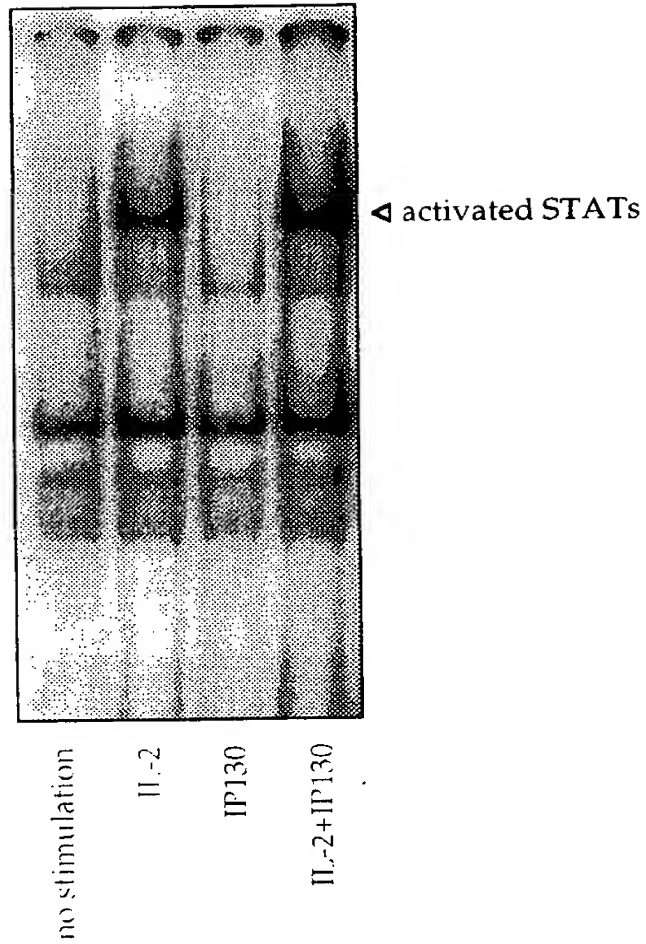
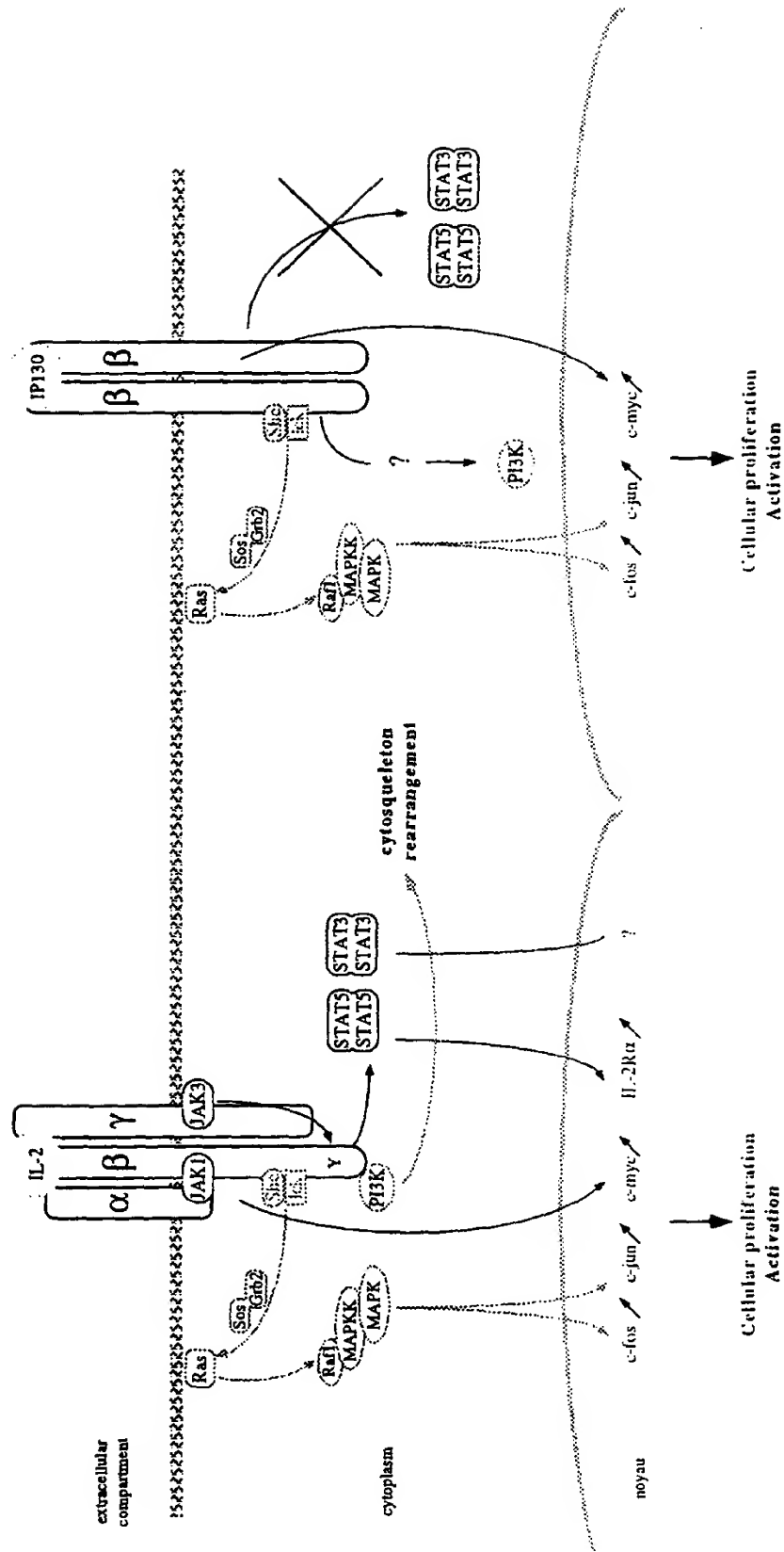


FIGURE 13



Model of transduction pathway induced by IP130

FIGURE 14

IL-2 receptor and its major signal transduction pathway

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NK cells (CD56⁺) entering in S+G2/M phases after IP130 stimulation
(synergy with IL-2)

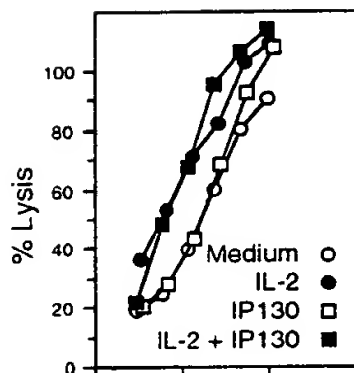
Treatment		J31	J32	J33
IL-2 50 nM		14	12	14
	IP130 60μM	0	17	≤5
	IP130 120μM	0	14	<5
IL-2 50 nM +	IP130 60μM	26	21	7
IL-2 50 nM +	IP130 120μM	28	28	28

FIGURE 15

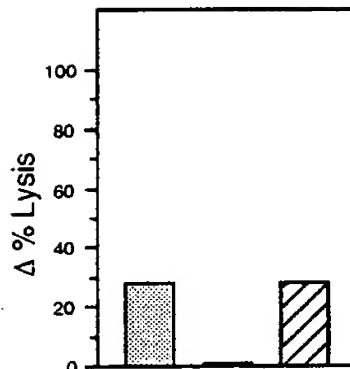
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Stimulation

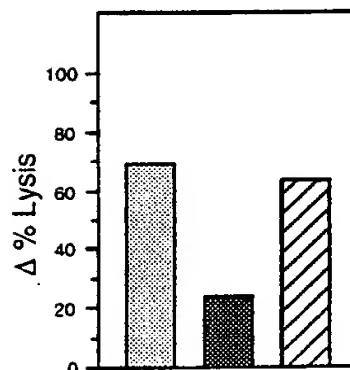
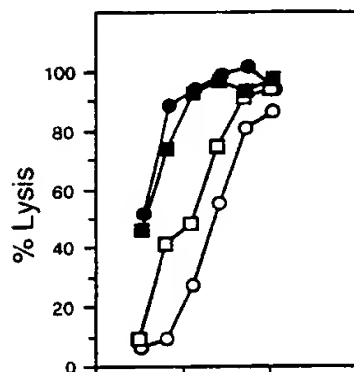
3 Days



Effector / target = 10



6 Days



9 Days

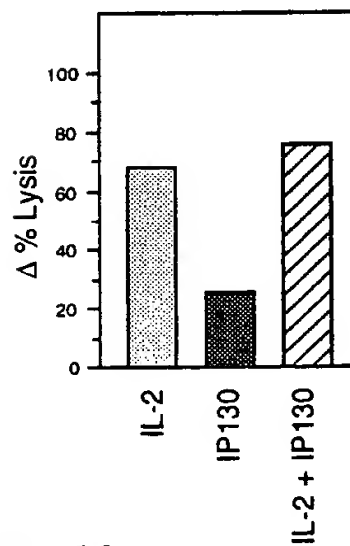
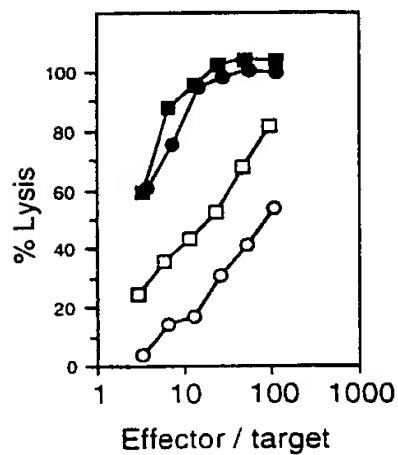


FIGURE 16